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#### IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

Applicants:

Baker et al.

Docket No:

39780-2830C1P9

Serial No:

10/006,485

Group Art Unit:

1647

Filed:

December 6, 2001

Examiner:

Rachel B. Kapust

For:

SECRETED AND TRANSMEMBRANE POLYPEPTIDES AND NUCLEIC

ACIDS ENCODING THE SAME

Commissioner for Patents Washington, D.C. 20231

# DECLARATION OF LUC DESNOYERS, Ph.D., DR. AUDREY GODDARD, Ph.D., DR. PAUL J. GODOWSKI, Ph.D., DR. AUSTIN GURNEY, Ph.D., DR. COLIN K. WATANABE and DR. WILLIAM WOOD, Ph.D. UNDER 37 CFR 1.131

- 1. We are the inventors of the above-identified application.
- 2. We have read and understood the claims pending in this application, and are aware that the claims have been rejected as anticipated by International Patent Application Publication No. WO 00/00610 (Lal *et al.*, publication date January 6, 2000).
- 3. We conceived and reduced to practice the invention claimed in the above-identified application in the United States prior to January 6, 2000.
- 4. At the time the present invention was made, one of the inventors, Luc Desnoyers, Ph.D., was, as still is, responsible for overseeing the testing of novel polypeptides, including the polypeptide designated PRO1412, in chondrocyte proliferation assay (Assay #111, Example 153). This assay is used to find agents that are capable of inducing chondrocyte proliferation and/or redifferentiation, and can, therefore, be used in the treatment of joint diseases using a tissue engineering approach or as promising drug candidates to repair aging or arthritic joints, for example, in which the chondrocytes have been dedifferentiated.

- 5. In this assay, isolated chondrocyte cells are seeded in 96 well plates with either serum-free medium (negative control), staurosporin (positive control) or the test PRO polypeptide. After 5 days, fluorescence dye is added to each plate and measured. The readout of the fluorescence from a plate containing the serum-free medium is measured to establish a background fluorescence level. A positive result in the assay is obtained when the fluorescence of the PRO polypeptide-treated sample is more like that of the positive control than the negative control. This type of fluorescence determination, wherein the readout is compared to positive and negative controls, is well known in the art.
- 6. A copy of a page from the Genengenes database which reports a positive result for the PRO1412 polypeptide encoded by DNA 64897-1628 (UNQ730) in Assay #111 is attached to this declaration (with its date redacted) as Exhibit A. The positive results reported in the database were also obtained prior to January 6, 2000.
- 7. Copies of pages from laboratory notebook showing the positive results for the PRO1412 polypeptide (SEQ ID NO:140), identified by Pin number PIN753-1, in Assay #111 are attached to this declaration (with dates redacted) as Exhibit B. These experiments were performed and the results were obtained prior to January 6, 2000.
- 8. Exhibits A and B clearly show that the polypeptide designated PRO1412 was tested, and its ability to induce the proliferation and/or redifferentiation was determined prior to January 6, 2000.

for Chin.	08/19/2004
Luc Desnoyers, Ph.D.	Date '
Audrey Goddard, Ph.D.	Date
Paul J. Godowski, Ph.D.	Date
Austin Gurney, Ph.D.	Date .
Colin K. Watanabe	Date
William Wood, Ph.D.	Date

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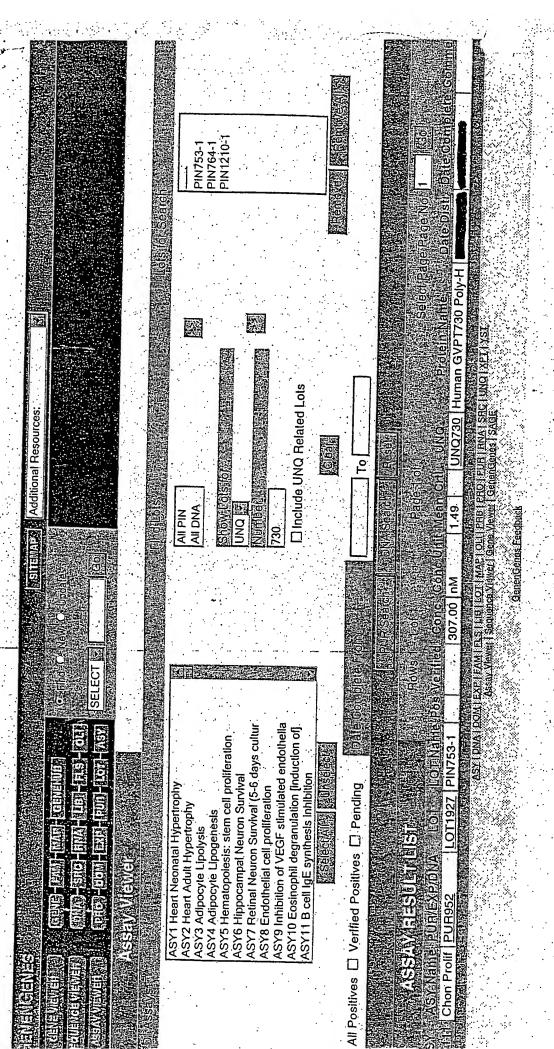
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Primary Assay Result Assay ID Assay Name Assay Date Chondrocytes Proliferation Assay

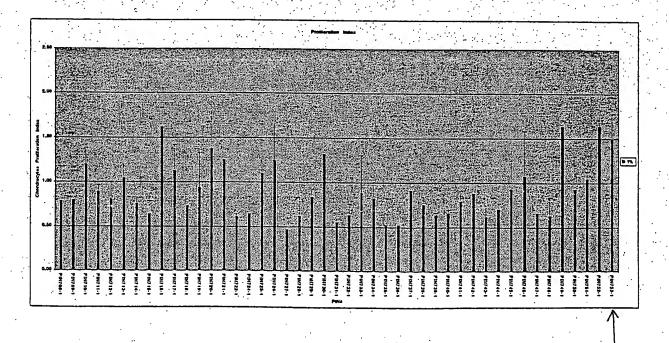
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		2	3	4	5	6	1 7			P1N742-1	PIN746-1	PIN750-1
	Sinurgsgonn	Staurosporin	Staurosporin	PIN717-1	PIN721-1	PIN725-1	PIN730-1	PIN734-1	PIN738-1	PINTAL-I	-141-10-1	
1 2	Media	Media	Media						PIN739-1	PIN743-1 .	PtN747-1	PIN751-1
1 .	PIN708-1	PIN711-1	PtN714-1	PIN718-1	PIN722-1	PIN727-1	PIN731-1	PIN735-1	Lat. 33m.	1 10,43	F	
ا آ							Dec 222 4	PIN736-1	P1N740-1	PIN744-1	PIN748-1	PIN752-1
	PIN709-1	PIN712-1	PIN715-1	PIN719-1	PIN724-1	PIN728-1	PIN732-1	PIN/ 3051		J		
					<u> </u>	- COLUMN 4	PIN733-1	PIN737-1	PIN741-1	PIN745-1	PIN749-1	PIN753-1
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4.0	- × A	55,1	67.1	98.4	100,4	173.2	166.5 103.6	166.8	103.1 75.5	89.3	104.3	78.5	119.8	]
- 1	18	81,1	-159.7 91.4	89.5	89.1	98.6 68.5	84.9	50.4	56.6	67.8	58.6	63.7	107.2· 119.6	1:
٠ [	. 6	54.5	102.4	75.6	70.8	55.9	37.7	59.7	47.5	74.3	72.1	52.5	175.3	1
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l	į.	115.0	125.9	159.3	105,5	77.5 161.4	57.8 124.0	64.6 128.4	73.5 125.7	95.4	137.9	172.1	. 139.5	] <

Conc			1.00%	<u> </u>	·		
PIN 4	N1	N2	AVERAGE	STOEV	Positive	Verified	Comments
PIN708-1	0.788	0,780	0,784	0.0	1		1
PIN709-1	0.949	0.545	0.797	0.2	1		1
PIN710-1	1.070	1.331	1,201	0.2	0 *	ľ	1 " - 1
PIN711-1	0.647	0.945	0.696	0.1		1	
PIN712-1	0,678	0.944	0.811	0.2		1	1967
PIN713-1	1.167	0.941	1.052	0.2			
PIN714-1	0.826	0.597	0,762	0.1			
PIN715-1	0.535	- 0.744	0.840	0.1			
PIN716-1	. 1.469	. 1.771	1,520	0.2	Positive		1
PIN717-1	0.925	1,333	1.129	0.3	1	1	
PIN718-1	0.622	0,653	0.738	0.1			
PIN719-1	.0.659	1,210	,0.938	0.4			
PIN720-1	0.973	1,786	1.380	0.0	Positive	1	3. 5
PIN721-1	1,598	. C.910 ·	1.254	0.5	4 .	1.	1 .
PIN722-1	0.632	0.597	0.614	0.0			1 ' '
PIN724-1	0.515	0.781	0.548	. 0.2	1 .		
PIN725-1	0.715	1,489	1.102	0.5	. ;		
PIN725-1	1,537	0.956	1.246	0.4	11		1 :
PIN727-1	0,599	0.343	0.471	0.2			
PIN728-1	0.471	0.774	0.623	0.2			. 1
PIN729-1	0.532	1,144	. 0.836	0.4		1 :	
PIN730-1	1.538	1,098	1.317	0.3	Positive .	1	·1 · · ·
PIN731-1	0.557	0.556	0.557	0.0			
PIN732-1	0.551	0.722	0.636	. 0.1		1	L
PIN733-1	0.595	1,184	0.890	0,4			1
PIN734-1	0.951	0,897	0.624	0,2		1	
PIN735-1	0.522	0,520	0.521	0.0	- I -	l .	
PIN736-1	0,438	0.617	0.527	0.1 .		1	
PIN732-1	0,678	1,159	0.919	0.3	i.		1 . 3
	0.586	0.824	0.755	0.1	1 .	1	.  -
PIN738-1 PIN738-1	0.624	0.654	0.639	0.0	1		1 .
	0.586	. 0,635	0,660	0.0	1		1
PIN740-1	0.712	0,860	0,796	0.1		1.	
PIN741-1	0,812	0.961	0.885	0.1	į.	1 .	
PIN742-1	0.541	0.695	0,618	0.1		1.:	
PIN743-1	0.665	0.751	0.708	Q.1	1.	[ ·	1
PIN744-1	0.599	1,272	0.935	. a.s	- 1	.   '	1 .
PIN745-1	1,436	0.724	1.060	0.5	1 -	j	1
PIN746-1	0.588	0,733	0.661	0.1	1 .	1.	
PIN747-1	0,388	0,781	0.633	0.2		1	
PIN748-1	1,684	1,565	1,636	0.1	Positive		
PIN749-1	0.757	1.105	0.931	0.2		.: <b>∤</b>	1 .
PIN750-1		1,104	1.046	0,1	1.	된 .	
PIN7\$1-1	0,989	1,665	1,842	0.0	Positive		
PIN752-1	1,616	1,287	1,491	0.3	Positive	1	
PIN753-1	1,695	1,207	1,481				

48	Average	STOEV
N708-1	0.78	0.0
N709-1 .	0.60	0.2
N710-1	1.20	0.2
N711-1	. 0.90	0.1
N7 12-1	0.81	0.2
V713-1	1.05	,0.2
N714-1	0.76	0.1
N715-1	- 0.64	. 0.1
N716-1	1,62	0.2
N717-1	1.13	0.3
N716-1	0.74	0.1
N719-1	0.94	0.4
N720-1	1,38	0.6
N721+1	1.25	0.5
N722-1	0.61	0.0
N724-1	. 0.65	0.2
N725-1	1.10	0.5
N726-1	1.25	. 0.4
N727-1	0.47	0.2
4728-1	0.62	0.2
V729-1	0.64	0.4
1730-1 ·	1.32	0.3
1731-1 -	0.56	0.0
1732-1	- 0.64	0.1
733-1	0.89	0.4
1734-1 ·	0.82	0.2
N735-1	0.52	0.0
1736-1	0.53	0.1
N737-1 .	0.92	. 0.3
N738-1	0.76	0.1
N739-1 N740-1	0.64	0.0
	0.60	0.0
N741-1	0.60	0.1
4742-1 4743-1	0.89	0.1
N/43-1 N/44-1	0.62	0.1
	0.71	0.1
N745-1 N746-1	6.94	0.5
4746-1 4747-1	1.08	- 0.5
	0.66	0.1
4748-1 4749-1	0.63	0.2
N749-1 N750-1	1.64	0.1
N730-1 N751-1	0.93	0.2
N752-1	1.05	0.1
N752-1	1.64 1.49	0.0



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